



**IN THE UNITED STATES PATENT AND TRADEMARK OFFICE**

Application No. : 10/524,060 Confirmation No: 4405  
Applicant : Zhiming SUO, M.D., Ph.D. and Barry W. FESTOFF, M.D.  
Filed : February 8, 2005  
Title : METHOD OF DETECTING AND PREVENTING  
ALZHEIMER'S DISEASE, PARTICULARLY AT  
PRODROMAL AND EARLY STAGES  
Group Art Unit : 1649  
Examiner : WANG, Chang Yu  
Atty. Docket No. : US 1421/05(VA)  
Date : October 11, 2005

**SECOND**  
**INFORMATION DISCLOSURE STATEMENT**

Commissioner for Patents  
P.O. Box 1450  
Alexandria, Virginia 22313-1450

Dear Sir:

Pursuant to 37 CFR 1.56 and 1.97-1.98, Applicants note herewith various references for the Examiner's consideration. These references include those cited on pages 35-44 of the application. (It is noted herewith that a copy of Reference No. 46, COWBURN et al., Receptor-G-protein signalling in Alzheimer's disease. Biochem. Soc.

Appl. No.: 10/524,060

Second Information Disclosure Statement dated October 11, 2005

Reply to Office Action of September 2, 2005

Symp., 2001, Vol 67, pp.163-175 was already submitted with the Information Disclosure Statement filed in the PTO on February 8, 2005.) The Examiner is respectfully requested to review and officially consider and make the reference(s) of record before issuing the first Office Action in connection with this case.

A completed Form 1449B/PTO listing the references thereon is enclosed herewith. (Pursuant to the PTO's Waiver signed by Deputy Commissioner Stephen G. Kunin on July 11, 2003, only the copy(ies) of foreign patent documents and non-patent literature, if applicable, is/are enclosed.) The Examiner is respectfully requested to return an initialed copy of Form1449B/PTO along with the next communication in connection with this case.

It is believed that no fee is due for this submission. Should that determination be incorrect, however, the Commissioner is hereby authorized to charge any deficiencies, or credit any overpayment, to our Deposit Account No. 01-0433, and notify the undersigned in due course.

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Should the Examiner have any questions or wish to discuss further this matter,  
please contact the undersigned at the telephone number provided below.

Respectfully submitted,

A handwritten signature in black ink, appearing to be 'DA' followed by a stylized flourish.

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**OTHER PRIOR ART - NON PATENT LITERATURE DOCUMENTS**

Examiner Initials <sup>1</sup>	Cite No. <sup>1</sup>	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the Item (book), magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), Publisher, country, where published, source.	T <sup>2</sup>
		Braak, H. & Braak, E. Neuropathological staging of Alzheimer-related changes. <i>Acta Neuropathol (Berl)</i> 82, 239-59 (1991).	
		Saitoh, T., Horsburgh, K. & Masliah, E. Hyperactivation of signal transduction systems in Alzheimer's disease. <i>Ann N Y Acad Sci</i> 695, 34-41 (1993).	
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		Sallese, M. et al. Regulation of G protein-coupled receptor kinase subtypes by calcium sensor proteins. <i>Biochim Biophys Acta</i> 1498, 112-21. (2000).	
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		Freeman, J.L., Pitcher, J.A., Li, X., Bennett, V. & Lefkowitz, R.J. alpha-Actinin is a potent regulator of G protein-coupled receptor kinase activity and substrate specificity in vitro. <i>FEBS Lett</i> 473, 280-4 (2000).	
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		Crawford, F. et al. Alzheimer's beta-amyloid vasoactivity: identification of a novel beta- amyloid conformational intermediate. <i>FEBS Lett</i> 436, 445-8 (1998).	

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		Tiruppathi, C. et al. G protein-coupled receptor kinase-5 regulates thrombin-activated signaling in endothelial cells. <i>Proc Natl Acad Sci U S A</i> 97, 7440-5 (2000).	
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		McLean C. A., R. A. Cherny, F. W. Fraser, S. J. Fuller, M. J. Smith, K. Beyreuther, Al Bush, and C. L. Masters. December 1999. Soluble pool of A $\beta$ amyloid as a determinant of severity of neurodegeneration in Alzheimer's disease. <i>Annals of Neurology</i> 46:860-866.	
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		Strittmatter W. J. February 2001. Apolipoprotein E and Alzheimer's disease: signal transduction mechanisms. <i>Biochemical Society Symposia</i> 67:101-109.	
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